Docket No. 00N-09390
Dockets Management Branch (HFA-305)
Food and Drug Administration
5630 Fishers Lane
rm. 1061
Rockville
MD 20852

September 22, 2000

Re: Self-nomination for Working Group on Markers of Drug-induced Vasculitis

I wish to nominate myself for the above Working Group, as announced in the July 26 issue of the Federal Register. I am a veterinary pathologist directing an ongoing research program into mechanisms of drug-induced vascular damage in rodents. Specifically, we have been addressing the possible roles of vasorelaxation, blood flow, and nitric oxide generation and have been using confocal laser microscopy to map the immunohistochemical expression of enzymes and receptors within the rat vascular wall. We have evaluated plasma von Willebrand factor and other molecules as potential markers of vascular damage and are extending our search for other surrogates using nuclear magnetic resonance spectroscopy of biofluids.

I believe that my experience and perspective in this area equip me well to contribute effectively. I confirm that I shall be willing to serve on the Working Group if selected.

Please find attached my biographical details plus a list of some more recent publications.

Sincerely,

Stephen J. Newsholme

Dr Stephen J. Newsholme, MMedVet, Diplomate ACVP, MRCVS
phone #: 610 - 270 - 7848

00N-0930

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Stephen John Newsholme

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Summary

A Preclinical Development scientist with a broad background in veterinary research together with more than 10 years experience as a toxicologic pathologist. Position as Senior Research Pathologist for a multinational pharmaceutical house has provided extensive experience in managing the preclinical safety assessment of assets from discovery through to product registration. Responsibilites have included representation on international project and discovery teams, leadership of multidisciplinary development teams and investigative programs and the mentoring of other scientists.

Positions Held – Past and Present

1989-	Currently Senior Research Pathologist, Safety Assessment, SmithKline Beecham Pharmaceuticals
1988-89	Assistant Director of Pathology, Smith, Kline & French Ltd., England
1985-88	Lecturer-Veterinary Pathology, University of Liverpool, England
1978-85	Senior Research Veterinarian – Pathology, Veterinary Research Institute, Onderstepoort, South Africa
1976-78	State Veterinarian, Natal, South Africa
1972-76	Veterinary practitioner, England

Professional Memberships

Diplomate, American College of Veterinary Pathologists
Member, Royal College of Veterinary Surgeons
Member, American Society for Investigative Pathology
Member, Society of Toxicologic Pathologists
Member, Society for Experimental Biology and Medicine
Associate Member, American Veterinary Medical Association

Personal Information

Nationality

American Citizen

Qualifications

1983 Diplomate, American College of Veterinary Pathologists
1982 MMedVet (Veterinary Pathology) cum laude, University of Pretoria
1972 BVetMed (Veterinary Medicine), University of London (equivalent to DVM)
1969 BSc (Physiology), University of London

Editorial Boards/Review Panels

Proceedings of the Society for Experimental Biology and Medicine, Toxicologic Pathology, Veterinary Pathology, Journal of Clinical Investigation

Recent Publications

Newsholme SJ, Thudium DT, Gossett KA, Watson ES, Schwartz L. Plasma von Willebrand factor as a biomarker for acute arterial damage in rats. (2000) *Tox. Pathol.* 28 (5).

Newsholme SJ, Maleeff B, Steiner S, Anderson NL, Schwartz L. Two-dimensional electrophoresis of liver proteins: characterization of a drug-induced hepatomegaly in rats. (2000) *Electrophoresis* 21:2122-2128.

Watson ES, Griswold DE, Schwartz W, **Newsholme SJ**. Theophylline induces splanchnic arterial damage in rats. (1999) *The Toxicologist 48 (1-S) Abst 1409.*

Webb EF, Tzimas MN, **Newsholme S**, Griswold DE. Intralesional cytokines in chronic oxazolone-induced contact sensitivity suggest roles for tumor necrosis factor α and interleukin-4. (1998) *J. Invest. Dermatol. 111:86-92.*

Invited Presentations

Newsholme SJ. Toxicology of PDE4 inhibitors. *Gordon Research Conference on Cyclic Nucleotide Phospodiesterases, Oxford, U.K., Sept 2000.*

Newsholme SJ. Safety of PDE4 inhibitors. Willian Harvey Research Conference, Nice, France, Dec.1999.
